

(19) World Intellectual Property
Organization
International Bureau



(43) International Publication Date
7 October 2004 (07.10.2004)

PCT

(10) International Publication Number
WO 2004/085465 A1

(51) International Patent Classification⁷: **C07K 7/00,**
C12Q 1/37

(21) International Application Number:
PCT/CA2004/000453

(22) International Filing Date: 25 March 2004 (25.03.2004)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
60/457,296 26 March 2003 (26.03.2003) US

(71) Applicant (for all designated States except US): **BIOMEPE INC.** [CA/CA]; 2901, rue Rachel Est, Suite 23, Montreal, Quebec H1W 4A4 (CA).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **BOILEAU, Guy** [CA/CA]; 7645 Malherbe, Brossard, Quebec H3C 3J7 (CA). **CARMONA, Adriana, Karaoglanovic** [BR/BR]; Alameda Sarutaia 103, Apto 161, CEP-01403-010 Sao Paulo, SP (BR). **CAMPOS, Marcelo** [BR/BR]; Rua

Rocha 46A, Apto 205, CEP-01330-000 Sao Paulo, SP (BR). **JULIANO, Maria, Aparecida** [BR/BR]; Rua Frei Gaspar 568, CEP-03048-020 Sao Paulo, SP (BR). **JULIANO, Luiz** [BR/BR]; Rua Rouxinol 795, Apto 94, CEP-04516-001 Sao Paulo, SP (BR).

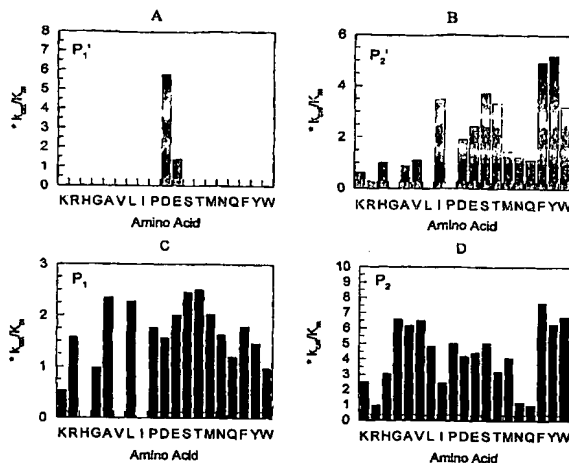
(74) Agents: **DUBUC, J. et al.**; Goudreau Gage Dubuc, Stock Exchange Tower, 800 Place Victoria, Suite 3400, P.O Box 242, Montreal, Quebec H4Z 1E9 (CA).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR,

[Continued on next page]

(54) Title: PHEX SUBSTRATES AND METHODS USING SAME



(57) Abstract: A fluorogenic PHEX substrate comprising a peptide unit; a fluorophore unit capable of conferring fluorescence on said substrate attached to an amino acid residue at a first end of the peptide unit; and a quencher unit capable of providing intramolecular quenching of said fluorescence attached to an amino acid residue at a second end of the peptide unit; the peptide unit having at least 6 amino acids residues including a sequence P₂-P₁-P₁-P₂ of 4 amino acid residues at positions P₂, P₁, P₁ and P₂ of the peptide unit, respectively; the amino acid residue at position P₂ being any amino acid residue; the amino acid residue at position P₁ being any amino acid residue except an isoleucine, a valine, or a histidine residue; the amino acid residue at position P₁ being an acidic amino acid residue selected from the group consisting of a glutamic acid residue and an aspartic acid residue, and being located at least 2 amino acid residues distal to both the fluorophore and the quencher units; the amino acid residue at position P₂ being any amino acid residue except a leucine, a proline or a glycine residue, with the proviso that said peptide unit does not have the sequence as set forth in SEQ ID NO:1. Methods of using the peptide sequence unit to identify PHEX modulators and for detecting PHEX in a sample.



GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK,
TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
ML, MR, NE, SN, TD, TG).

— *before the expiration of the time limit for amending the
claims and to be republished in the event of receipt of
amendments*

Published:

— *with international search report*

*For two-letter codes and other abbreviations, refer to the "Guid-
ance Notes on Codes and Abbreviations" appearing at the begin-
ning of each regular issue of the PCT Gazette.*